

Review article

Prevalence of chronic wounds in the general population: systematic review and meta-analysis of observational studies



Laura Martinengo, MD ^a, Maja Olsson, RN, MPH ^a, Ram Bajpai, PhD ^a,
Michael Soljak, MD, PhD ^a, Zee Upton, PhD ^{b,c}, Artur Schmidtchen, MD, PhD ^{c,d,e},
Josip Car, MD, PhD ^{a,f,*}, Krister Järbrink, PhD ^a

^a Centre for Population Health Sciences (CePhaS), Lee Kong Chian School of Medicine, Nanyang Technological University, Singapore, Singapore

^b Institute of Medical Biology, Agency for Science, Technology and Research (A*STAR), Singapore, Singapore

^c Dermatology, Lee Kong Chian School of Medicine, Nanyang Technological University, Singapore, Singapore

^d Division of Dermatology, Department of Clinical Sciences, Biomedical Center B14, Lund University, Lund, Sweden

^e Copenhagen Wound Healing Center, Bispebjerg Hospital, Department of Biomedical Sciences, University of Copenhagen, Copenhagen, Denmark

^f Global eHealth Unit, Department of Primary Care and Public Health, School of Public Health, Imperial College London, London, UK

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ABSTRACT

Purpose: Chronic wounds are a major public health challenge, but little is known about the true burden with studies reporting different estimates because of disparities in study designs and measurement methods. This hampers efficient resource allocation, planning, and improvement of wound care.

Methods: Our study aimed to pool prevalence estimates from a global perspective by systematically carrying out searches in MEDLINE, EMBASE, Cochrane, CINAHL, Global Health, and PsycINFO databases for articles reporting the prevalence of chronic wounds in adults, from January 2000 to June 2018. The included publications had to define wound chronicity by duration (≥ 3 weeks), and/or labeling the wounds as chronic, complex, or hard-to-heal.

Results: Seventeen studies met the inclusion criteria, and 11 studies analyzing chronic wounds in the general population were included in random effects meta-analyses to calculate pooled prevalence. Chronic wounds of mixed etiologies ($n = 3$) showed a pooled prevalence of 2.21 per 1000 population, and for chronic leg ulcers ($n = 9$), the prevalence was estimated at 1.51 per 1000 population.

Conclusions: Our findings, aligned to previous studies reporting point prevalence of chronic wounds identified within the healthcare system, showed that the vast majority of chronic wounds in epidemiological studies are made up by chronic leg ulcers.

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Introduction

Skin diseases are a growing problem worldwide. The global burden for skin and subcutaneous diseases has increased rapidly over a 10-year period, with a prevalence of 605,036,000 in year 2015 compared to 492,883,000 in 2005 based on global, regional, and national data from over 195 countries and territories reported by the Global Burden of Disease (GBD) study [1]. Even though there

are subcategories within the broad “skin and subcutaneous diseases,” specific prevalence data for chronic wounds are inadequate, and seldom presented separately [1]; hence, no global prevalence data are available for chronic wounds. Estimating wound prevalence is challenging, as there is no agreement or recognized distinction between acute and chronic wounds or a clear pre-established consensus for chronicity [2]. A commonly used definition, labeling chronic wounds as wounds that fail “to proceed through an orderly and timely process to produce anatomic and functional integrity” [3], does not specify any timeframe for the healing or at what time point the wound becomes chronic. This absence of a coherent and systematic classification introduces difficulties in calculating reliable pooled prevalence estimates unambiguously for chronic wounds in the general population across studies and hinders improvement efforts.

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* Corresponding author. Associate Professor Josip Car, Centre for Population Health Sciences (CePhaS), Lee Kong Chian School of Medicine, Nanyang Technological University, Clinical Sciences Building 11 Mandalay Road, Singapore 308232, Singapore.

E-mail address: josip.car@ntu.edu.sg (J. Car).

Although there are several studies from different levels of care (e.g., hospitals, community care, and nursing homes) from countries across the world [4–7], wounds are in many aspects an under-reported health issue [8]; despite the fact that they are associated with significant costs [9–11], they have become a major challenge to healthcare systems worldwide [12]. To have accurate prevalence estimates for chronic wounds, it is fundamental to understand the disease burden, inform decision makers [13,14], and improve planning and the delivery of health services [14]. For example, a high prevalence of chronic wounds will demand more nurses with wound-care expertise, or it could aid in allocating resources to target the most pressing health problems in the area [13,14]. Prevalence estimates for chronic wounds could potentially make healthcare service provision more cost-efficient when incorporated in health economic evaluations exploring how to reduce costs [15].

To accurately assess the prevalence of chronic wounds is problematic because of disparities in study design and how health conditions are defined and measured [16]. Factors that limit comparability of prevalence studies are heterogeneous study design and lack of clear definitions [14]. These obstacles hamper accurate comparability and calculations of pooled estimates for chronic wounds. This is likely a plausible reason why to date there have been no comprehensive systematic review with a meta-analysis exploring the global magnitude of the condition. The aim of this systematic review is to make an attempt to pool a prevalence estimate for chronic wounds globally within the general population based on existing literature and by applying a clear definition of chronicity (duration and/or wounds labeled as chronic).

Material and methods

Search strategies

This systematic review followed the Preferred Reporting Items for Systematic Review and Meta-Analysis (PRISMA) guidelines [17]. The protocol was published and registered with PROSPERO (CRD42016037355) [18]. The search strategy was developed by the research team in collaboration with experienced medical research librarians at Lee Kong Chian School of Medicine and Karolinska Institutet. An initial search was carried out in March 2016 covering MEDLINE (Ovid), EMBASE (Ovid), EBM Reviews and Cochrane Database of Systematic reviews (Ovid), Cumulative Index to Nursing and Allied Health Literature (CINAHL), (EBSCO), PsycINFO (EBSCO), and Global Health (EBSCO) databases for articles published from January 2000 through December 2015. An additional search was also performed in August 2016 to incorporate the newly developed term “pressure injury”. In June 2018, we updated the search to include studies published from January 2016, using the original strategy with the addition of “pressure injury”. The reference lists of all included articles were also searched for other sources of information [18].

For this review, chronic wounds were defined as those with a duration of 3 weeks or longer or if the wound was referred to as chronic, hard-to-heal, or complex, even when there was no mentioning of duration. This definition was deemed appropriate as there is no pre-established consensus for chronicity [2], and previous literature has suggested that wounds that do not reduce size after 2 to 4 weeks are likely to become chronic [19–21]. This rather generous cut-off for duration allowed us to comprehensively capture a substantial amount of publications presenting prevalence data for chronic wounds.

Inclusion and exclusion criteria

All included articles had to present prevalence estimates of chronic wounds in adults within the general population. We

included articles referring to chronic wounds in the categories of venous ulcers, arterial ulcers, diabetic ulcers, and pressure ulcers. Studies had to be of a cross-sectional or cohort design, published in peer-reviewed journals, in English and available as full text publications. We excluded studies on chronic wounds of other etiologies, for example, injuries due to trauma, infections, autoimmune diseases, surgical wounds, and skin tumors that due to size and malignancy require a longer time to heal. Randomized controlled trials, case studies, reviews, editorials, and abstracts were excluded, as well as studies that presented prevalence estimates of specific subsets of populations such as the terminally ill, patients with spinal cord injuries, or patients in special care units.

Study selection, data extraction, and quality assessment

Two reviewers screened titles and abstracts independently to identify studies for potential inclusion. The full text of these articles was retrieved and reviewed by two investigators to ensure compliance with the eligibility criteria. Studies excluded in this step were listed with the reason for their exclusion. We also searched the bibliographic references of all included articles for additional studies. The selection process for articles was summarized in a flow chart. Information about study details (e.g., title, journal, author, year, country), study design (type of study, aims, method for data collection, response rate, recruitment, sample methods, and inclusion/exclusion criteria), participants' characteristics (number of persons surveyed, population characteristics, i.e., setting, age, gender, ethnicity, demographics), specific wound data and complications (diagnostic methods, ulcer specifications including stage), and prevalence, or sufficient information to calculate an estimate of the prevalence was extracted for each article by two reviewers. This information was collected in an electronic data extraction form that was previously piloted and reviewed by a panel of systematic reviews experts. The included articles were quality assessed using the Joanna Briggs Institute (JBI) critical appraisal checklist for studies reporting prevalence data [22] by two reviewers. The tool evaluated the following: sample frame, sampling methods, sample size, description of setting, sufficient coverage for data analysis, objective and reliable methods for identifying the condition, appropriate statistical analysis, and adequate response rate. The tool appraises each domain as Yes/No/Unclear/Not applicable. We assigned a value 1 to a Yes answer and a value 0 if the answer was No/Unclear/Not Applicable. A higher score denotes a higher quality study, with a maximum attainable score of 9. As the appraisal tool does not give a specific definition of an acceptable quality, we defined as acceptable quality scores equal to or higher than five. In all stages of this process, discrepancies between reviewers were resolved by consensus or, if not, resolved by arbitration by a third reviewer.

Data synthesis and statistical analyses

Prevalence was calculated for all chronic wounds regardless of the etiology and for the leg ulcers subgroup. Prevalence was calculated as the ratio between the total number of reported ulcer cases over the total study population and presented as the number of cases per 1000 population. Random-effects meta-analysis, using DerSimonian and Laird method [23], was employed on individual study estimates to obtain a crude summary estimate for prevalence. Standard errors and 95% confidence intervals (CI) were determined from the reported crude estimates and population denominators, assuming exact binomial distribution as described by Clopper and Pearson [24]. Heterogeneity between studies was assessed using the χ^2 -based Cochran's Q statistic and I-squared (I^2) statistics (>50% representing moderate heterogeneity) [25], and a subgroup analysis was conducted to identify potential sources of heterogeneity. A

prediction interval for the random-effects distribution was also calculated to understand the possible range of ulcer prevalence if a new study is conducted as suggested by Higgins and Thompson [26]. Publication bias was assessed by funnel plot and its asymmetry was tested by Egger's linear regression method [27] ($P < .10$ was considered as significant). The influence of studies on overall results was graphically assessed by Baujat's plot [28]. Because of demographic and epidemiological transitions, it is understandable that the prevalence rates of ulcers may vary with the mean age of patients and the time the study was carried out. Therefore, with the limited data, a univariate and multivariate meta-regression analysis was conducted using maximum likelihood method for age and year of study ($P < .10$ considered significant given the low power of these tests). A P value of $< .05$ was considered statistically significant for the effect of study-level covariates on the estimated prevalence [29]. All statistical analyses were conducted on R software (Version 3.3.2) using "meta" package [30].

Results

The systematic search returned a total of 5805 publications after excluding duplicates. A subsequent search for "pressure injury" returned a total of 213 unique references, producing an overall total of 6018 articles. After excluding irrelevant articles based on titles and abstracts, 716 articles remained for full text examination. Unrelated articles not fulfilling our inclusion criteria

were discarded, and two articles from manual searches of the included articles reference lists were added, summing up to 15 remaining articles (Fig. 1). The updated search in year 2018 yielded yet another two articles, for a final grand total of 17 articles included in the study.

Quality assessment

The quality assessment indicated that most included studies were of acceptable quality. The average score was 6.6 points (range 5–9, with higher scores denoting higher quality articles), where the best scores were associated with sampling, sample frame, methods for sampling, and sample size. All studies used appropriate sampling strategies for the targeted populations (i.e., surveyed healthcare settings that were probable to treat wound patients), presented sufficient sample sizes, and most articles described the study settings in detail. The lowest scores were obtained in the questions referring to methods to assess the condition (correct diagnosis and staging of chronic wounds) and the appropriateness of the statistical analysis used. A summary of the quality assessment is included as [Additional File 1](#).

Characteristics of included studies

Seventeen studies conducted in 13 different countries reported prevalence of chronic wounds of various etiologies. The included

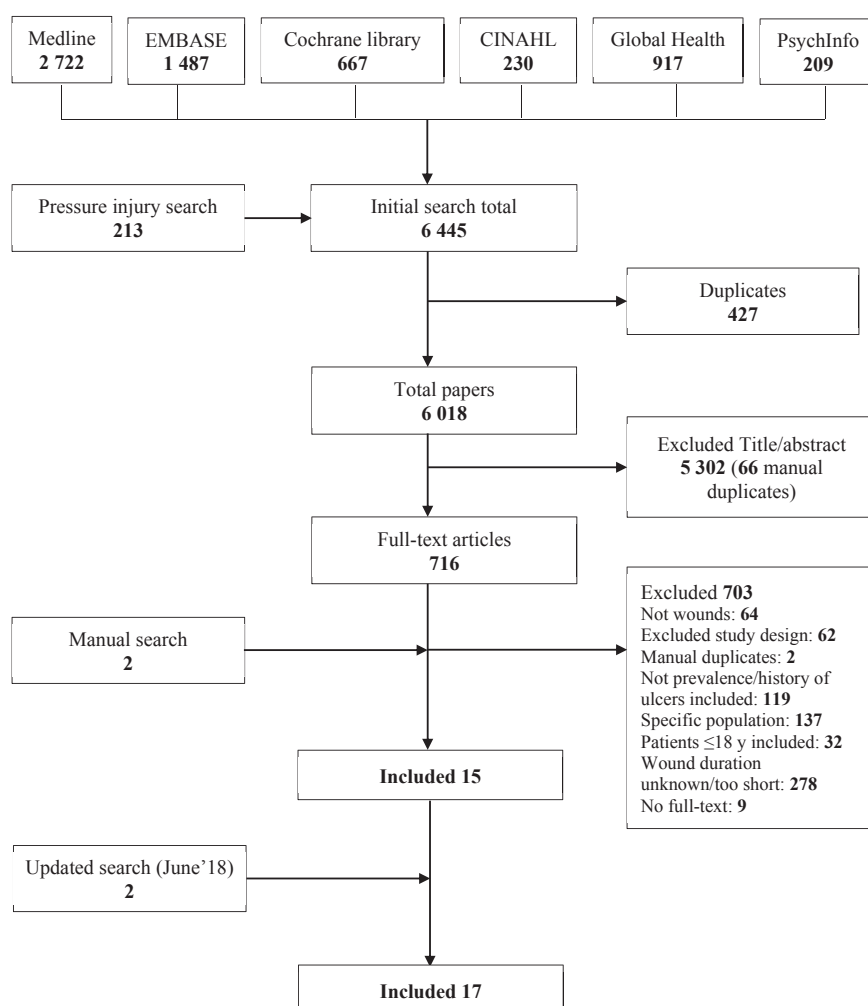


Fig. 1. Flow chart to illustrate the selection process.

studies were published between 2000 and 2017, and the majority were from European countries (65%). When the studies reported it, the patients' mean age ranged from 50 to 80 years of age. All studies reported prevalence for chronic wounds obtained during a limited period ranging from 1 day to 2 years. The total population covered across studies was 15,748,658 persons, of which 3,503,756 were patients in subgroup populations, for example, hospitalized patients only, etc. The total number of chronic wound cases was 141,311 (46,000 in subgroup populations). Most included patients ($n = 9,109,732$) and wounds ($n = 91,723$) were made up from one study from Germany assessing chronic wounds in the general population [31]. The most commonly reported type of chronic wounds were leg ulcers ($n = 9$) reported in 53% of the studies [31–39]. Five studies reported chronic wounds of various etiologies [40–43], three studies reported pressure ulcers [44–46], and one study assessed diabetic foot ulcers [47]. Characteristics of the included studies are presented in Tables 1 and 2.

In the studies presenting leg ulcers, the main reported underlying etiology was venous disease [32–38,40]. However, one article investigating hospital settings reported diabetes as the main etiology for leg ulcers [42]. Another study assessing leg ulcers specified foot ulcers as a subgroup and stated diabetes as the most common origin for this group [35]. Four studies presented various types of chronic wounds, for example, leg ulcers, pressure ulcers, diabetic foot ulcers, and mixed chronic ulcers [31,40,42,43].

Six studies presented chronic wounds prevalence solely from specific settings: patients in home care [43,44], hospitals [42,45,46], and a hospital-based diabetes clinic [47].

Prevalence stratified by sex and age

All included studies except for three [31,44,46] presented chronic wound prevalence estimates stratified by sex. Chronic wounds were more prevalent in women (range: 53%–68%), except in one hospital-based study from China where the proportion of chronic wounds was higher among men (64%) [42].

Regarding age, most studies reported rather similar mean/median ages (around 70–80 years of age), with the exception of one study from Egypt on prevalence of diabetic foot ulcers that reported a mean age of 50 years [47].

Meta-analyses for chronic wounds

Eleven studies were included in the meta-analysis. All included studies presented prevalence data in the general population [31–41]. The test for consistency across studies revealed substantial heterogeneity ($I^2 = 100\%$; $P < .01$). The meta-analysis estimated a total prevalence of chronic wounds of 1.67 per 1000 population (C.I.: 0.83–2.80). Subgroup analyses based on the type of wounds revealed a combined prevalence of 2.21 per 1000 population (C.I.: 0.56–4.94) for studies reporting ulcers of more than one etiology [31,40,41]. The meta-analysis of studies reporting prevalence of leg ulcers [31–39] computed a prevalence of 1.51 per 1000 population (C.I.: 0.24–3.84) (Fig. 2). The I^2 value for leg ulcer subgroup was found to be 100% ($P < .01$), indicating the presence of substantial heterogeneity between studies. The estimated prevalence between chronic ulcers of different etiologies and leg ulcers was not

Table 1
Characteristics of studies included in the meta-analysis

| Author (y) | Country | Wound type | Age (mean) | Total population | No. of wound cases | Notes (time of data collection) | Reported prevalence |
|----------------------------------|----------------|--|---|------------------|--------------------|---|---------------------|
| Clarke-Moloney et al 2006 [32] | Ireland | Leg ulcers (>1 mo) | 75.5 | 339,591 | 382 | Survey to nurses, patient seen (1 wk) | 0.11% |
| Hall et al 2014 [40] | United Kingdom | Complex wounds defined as “superficial-, partial- or full-thickness skin loss healing by secondary intention” | ~70* | 751,485 | 1,093† | Ascertainment by healthcare providers (2 wk) | 0.15% |
| Heyer et al 2016 [31] | Germany | Chronic wounds defined by chronic wound diagnosis of any wound etiology (ICD-10). Stratified by ulcer types, diabetic foot ulcer, pressure ulcer, and leg ulcer. | Mean/median not reported. Range: 20–>90 | 9,109,732 | 91,723 | Register data from patients insured with the nationwide operating statutory health insurance provider (2 y) | 1% |
| McDermott-Scales et al 2009 [41] | Ireland | Chronic open leg ulcers of varying etiology located below the knee on the leg or foot (>4 wk), and other wounds or skin injury resulting in tissue damage | Mean: 69.3 | 133,562 | 179 | Survey to nurses in community care settings. (1 random day) | 0.13% |
| Moffatt et al 2004 [33] | United Kingdom | Open wounds on the leg excluding foot ulcers (>4 wk) | 75 | 252,000 | 113 | Ascertainment by healthcare professionals NHS (4 wk) | 0.04% |
| O'Brien et al 2000 [34] | Ireland | Open sore anywhere below the knee (>4 wk) | 72.3 | 317,069 | 374 | Ascertainment by healthcare professionals (2 mo) | 0.12% |
| Oien et al 2000 [35] | Sweden | Chronic leg and foot ulcers | 79‡ | 151,610 | 316 | Ascertainment by community and district nurses (1 wk) | 0.19% |
| Oien et al 2006 [36] | Sweden | Leg and foot ulcers (hard-to-heal) | 81‡ | 150,334 | 229 | Ascertainment by community and district nurses (1 wk) | 0.15% |
| Palsdottir et al 2010 [37] | Iceland | All ulcers below the knee (≥ 6 wk) | 75.2 | 313,376 | 226 | Ascertainment by healthcare (2 wk) | 0.07% |
| Pina et al 2005 [38] | Portugal | Open wounds on the leg (>4 wk) | 70.2 | 185,708 | 250 | Ascertainment to health centers (2 wk) | 0.13% |
| Walker et al 2002 [39] | New Zealand | Breaks in the skin on the lower leg (below the knee) or on the foot (>6 wk) | 75‡ | 540,435 | 426 | Ascertainment by community and hospital-based health professionals or self-notification (1 y) | 0.08% |

* Mean of the total number of ulcers (including nonchronic).

† 1093 refers to ulcers in people older than 18 years of age (contact with authors).

‡ Age reported as median.

Table 2
Characteristics of prevalence studies for chronic wounds in specific settings

| Author (y) | Country | Type of wound | Age (mean) | Total population | Sub population | No. of wound cases | Notes | Setting | Reported prevalence |
|---------------------------|---------------|--|----------------------------|------------------|--------------------------------|--------------------|--|---------------------------------------|---------------------|
| Amir et al 2017 [46] | Indonesia | Pressure ulcers (>3 mo) | 55.6 | — | No. of patients 1132 | 8 | Hospitals (1 d) | Only hospitals | 0.70% |
| Barrois et al 2008 [45] | France | Pressure ulcers (>1 mo) | Females 72.3 Males 70.1 | — | No. of patients: 37,307 | 1671 | Both chronic and nonchronic | Nonuniversity hospitals | 4.48% |
| El-Nahas et al 2008 [47] | Egypt | Diabetic foot ulcers (>4 mo) | 50.47 | — | 1220 | 15 | Outpatient diabetes clinic (over 1 y) | Specialist diabetes outpatient clinic | 1.2% |
| Jiang et al 2011* [42] | China | A chronic wound was defined as a defect in the skin for example, pressure, diabetic, and venous wounds (>1 mo) | — | — | 1,488,201 | 2513 | Hospitals (2 y) | Only hospitals | 1.69% |
| Rodrigues et al 2006 [43] | Canada | Chronic wounds presented as pressure ulcers, venous ulcers, and diabetic foot wounds. | 68.5 | — | 34,857 (patients in home care) | 488 | Ascertainment by community and home care nurses (2 mo) | Patients in home care | 1.4% |
| Schwien et al 2005* [44] | United States | Pressure ulcers (nonhealing) | — | 1,941,039 | — | 41,305 | Database home care setting | Home care setting | 2.13% |

* Age not reported.

statistically significant ($P = .625$). The prediction interval (C.I.: 0.00–8.16 per 1000 population) showed the confidence band of estimated prevalence for future population-based observational studies on chronic wounds. The weights of the studies reported from random effect model ranged from 25% for studies with ulcers of more than one etiology, and 75% for studies with leg ulcers.

Meta-regression analysis

The overall association between the pooled prevalence of chronic wounds, mean reported age (years), and year of reporting was assessed by univariate and multivariate mixed-effect meta-regression analysis. The estimated effect of age and reporting year was not found to be statistically significant in univariate (age, $P = .102$; year of reporting, $P = .449$) and in multivariate regression

analyses (age, $P = .156$; reporting year, $P = .814$) (see [Supplementary Figs. 1 and 2](#) for bubble plots). The overall analysis indicated a negligible impact of age and year of reporting on the estimated prevalence of chronic wounds in general population.

Publication bias

We used funnel plots ([Fig. 3](#)) to assess publication bias. In the figure, the vertical line represents the summary estimate of the chronic wound prevalence. The diagonal lines represent the 95% confidence limits around the summary treatment effect. These show the expected distribution of studies in the absence of heterogeneity or selection biases. The funnel plot was asymmetric and indicated there might be missing studies. The funnel plot asymmetry was assessed by Egger's linear regression test that showed

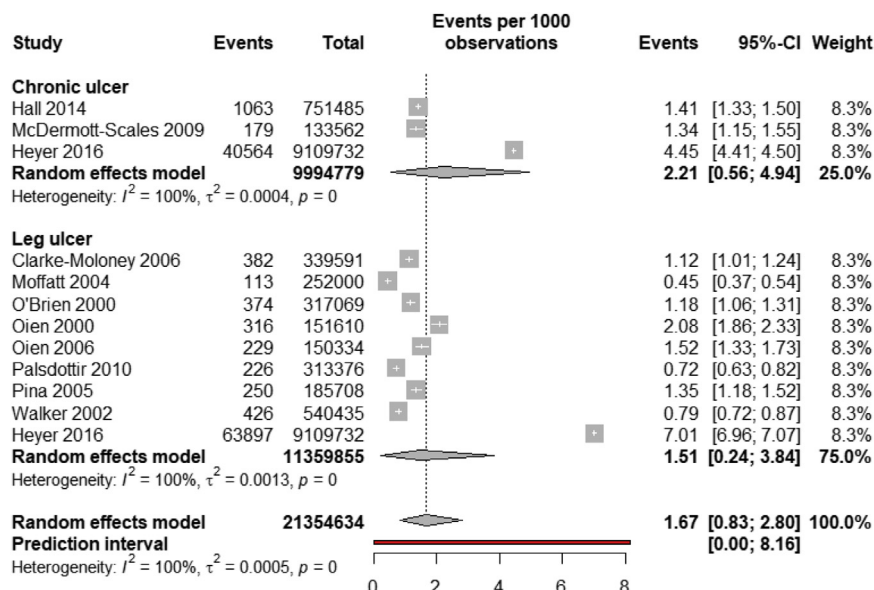


Fig. 2. Forest plot (pooled prevalence of chronic and leg ulcers per 1000 population).

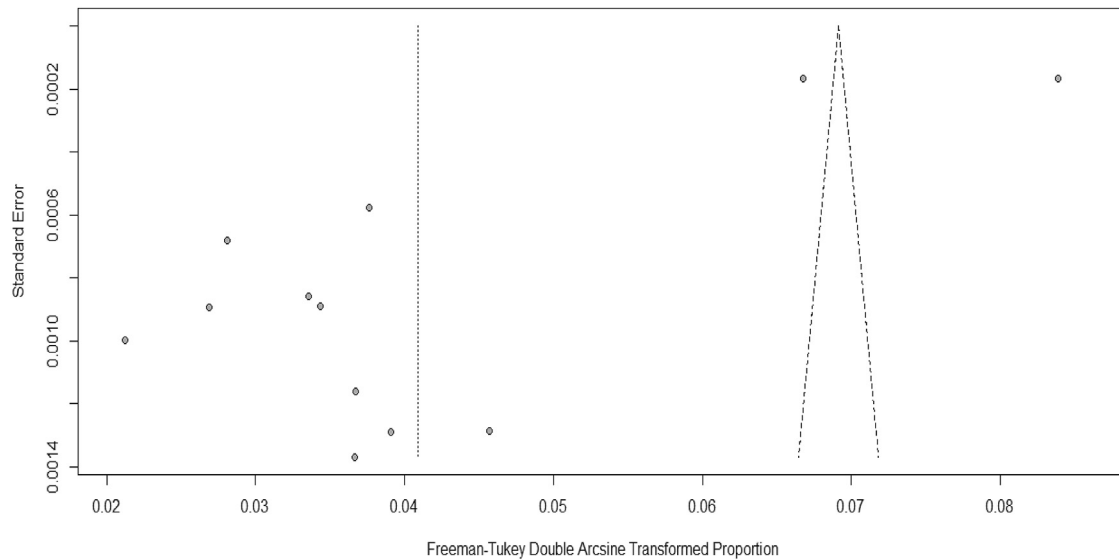


Fig. 3. Funnel plot with 95% confidence limits showing the prevalence (p) of ulcers in each study by the standard error (SE) of the studies.

significant publication bias (slope = 0.08, $t = -4.38$, $P < .01$). The overall influence of included studies on heterogeneity was assessed by Baujat plot, which indicated half of the studies were the larger contributors of the reported heterogeneity ([Supplementary Fig. 3](#)).

Discussion

This study, applying a clear definition for chronicity, provides a conservative estimate of the prevalence of chronic wounds from a global perspective.

Eleven of the 17 included studies presented chronic wounds prevalence in the general population. Of these, two studies included ulcers of various etiologies, eight studies reported leg ulcers, and one study presented the prevalence of chronic wounds of various etiologies and leg ulcers. These studies were incorporated in meta-analyses. Ulcers of various etiologies had a pooled prevalence of 2.21 per 1000 population; however, because of the small number of included studies, this result should be considered with caution. The pooled prevalence of nine studies reporting leg ulcers was estimated at 1.51 per 1000 population. Our results are generally aligned to previous studies reporting point prevalence of chronic wounds identified within the healthcare system [48,49], although they differ from studies reporting lifetime prevalence of chronic ulcers that tend to be approximately 10 per 1000 population [48,50]. Given that all included studies presented prevalence of wounds detected within the healthcare system, self-treated cases were not reported.

Although we aimed to estimate the global prevalence of chronic wounds, our study mainly consisted of publications from developed countries (82%), with only three articles from developing countries. These articles reported setting-specific prevalence of chronic wounds [42,46,47], and none were included in our meta-analysis. The dearth of epidemiology data from low-income countries was also revealed in a previous prevalence review for chronic wounds by Graves et al. [7], which presented only two papers from developing countries out of 69 included articles.

Global prevalence estimates of chronic wounds, including high-quality studies from developing countries, are urgently needed to understand geographical and ethnic variations in prevalence [51], as well as the leading etiology in each region. Previous reports from developing countries in the tropics show

that chronic wounds are usually secondary to chronic infectious diseases like leprosy or tuberculosis [52] and burns [6], but with the rising prevalence of diabetes, the prevalence of chronic wounds is also increasing [53]. This information can be used as guidance when planning for healthcare services and provision of treatments globally [54]. Obtaining wound data on a population level in developing countries is, however, problematic because of several limitations, for example, lack of healthcare information systems, limited access to adequate documentation records [55], workforce shortages impeding detailed documentation, and the ability to participate in surveys about prevalence [56]. In some developing countries, epidemiology data for chronic wounds have been collected in hospitals [6,57], but these appear insufficient to estimate the prevalence in the general population [57]. To estimate the global prevalence of chronic wounds accurately, future studies must obtain comprehensive data from developing countries as well as developed ones. Accurate prevalence data on chronic wounds is currently deficient even though Global Burden of Disease studies have assessed the burden of skin diseases quite comprehensively [58,59]. In regard to chronic wounds, studies fail to address them as a separate category, and presumably include them in the subgroup “Other skin and subcutaneous diseases,” with the exception of pressure (decubitus) ulcers that are presented as a standalone subgroup [58,59]. Hay et al. [59] in their study on the global burden of skin disease in 2010 included data from 187 countries and showed that “Other skin and subcutaneous diseases” was the fifth most prevalent disease globally. There is no description of the skin diseases that populate this group or of the relative contribution to each group in the final calculation of prevalence. The data on decubitus ulcers showed highly variable prevalence rates across regions, fluctuating from 0.168 cases per 1000 population in Central American females to 2.324 cases per 1000 population in North African and Middle Eastern females [59]. This shows the magnitude of skin diseases but also confirms the lack of specific epidemiology data for chronic wounds, leaving a huge gap in global skin research. The complexities associated with a lack of clear diagnosis and uniform methods for measuring and categorizing chronic wounds (e.g., diverse duration thresholds for wound chronicity) make it challenging to accurately explore and estimate the condition across countries, often resulting in under-reporting [51].

However, in exploring the global burden of chronic wounds, it is urgent to assess prevalence in both developed and developing countries [51,54].

Regarding study quality, none of the included studies employed a random sampling strategy. All prevalence cases included were sampled from patients already within the healthcare system so under-reporting is likely. Although chronic wounds are often cared for within the healthcare system, efforts should be made to capture those patients caring for their wounds themselves. Other strategies, including door-to-door surveys, could be considered to capture all patients with a chronic wound, within and outside the healthcare system, for example, in rural areas [52]. Furthermore, no study reported the differences between characteristics for included and nonincluded participants; this could also introduce a nonresponse bias.

This study is not without limitations. First, although we tried to develop a comprehensive search strategy including all ulcer categories, and as many variations of the terminology, that included running a supplementary search when informed of the newly incorporated term “pressure injury,” we may have inadvertently missed relevant studies.

Second, this systematic review and meta-analysis attempted to pool estimates for global chronic wound prevalence observed a large statistical variability (I^2 value). Examining the heterogeneity of included studies is a crucial part of a meta-analysis as observational studies are prone to biases because of various clinical and methodological designs such as inclusion criteria, definition of chronicity, and sampling strategies. The result of a meta-analysis should therefore be viewed as an approximation of the actual prevalence and be interpreted with caution. However, statistical heterogeneity due to differences in methods and/or assessment or definitions of chronic wounds across studies suggests the use of different measurement methods but do not necessarily indicate that the true prevalence varies. In this sense, the prediction interval can be helpful to map the range of actual prevalence in future studies. Sources of heterogeneity in this review could likely be linked to the use of nonstandardized methods across studies and various procedures for data collection. Third, the diversity of assessment methods for chronic wounds makes it difficult to capture prevalence data across studies and countries, hence impedes a robust global estimate [48]. Various definitions for chronicity (a couple of weeks up to several months) and the lack of standardized diagnostic methods, for example, leg ulcers, should ideally be confirmed by noninvasive methods such as Doppler, which introduces biases leading to either underestimations or overestimations of the existence of chronic wounds [48,60]. This becomes problematic when ulcers are multifactorial or when prevalence studies present leg ulcers of various etiologies (e.g., diabetic foot ulcers and leg ulcers) clustered together and presented as leg ulcers [50]. Reliable and accurate prevalence data for chronic wounds according to etiologies are of paramount importance when planning for proper treatment because different underlying causes of wounds demand different treatments and intervention strategies [60].

To conclude, our findings showed that the vast majority of chronic wounds are made up by chronic leg ulcers. However, prevalence studies with clear definitions of chronicity and comparable estimates are urgently needed to understand the magnitude of chronic wounds and to globally assess their prevalence.

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Supplementary data

Supplementary data associated with this article can be found in the online version at <https://doi.org/10.1016/j.annepidem.2018.10.005>.

References

- [1] Vos T, Allen C, Arora M, Barber RM, Bhutta ZA, Brown A, et al. Global, regional, and national incidence, prevalence, and years lived with disability for 310 diseases and injuries, 1990–2015: a systematic analysis for the Global Burden of Disease Study 2015. *Lancet* 2016;388(10053):1545–602.
- [2] Kyaw BM, Jarbrink K, Martinengo L, Car J, Harding K, Schmidtchen A. Need for improved definition of “chronic wounds” in clinical studies. *Acta Derm Venereol* 2018;98(1):157–8.
- [3] Lazarus G, Cooper D, Knighton D, Margolis D, Pecoraro R, Rodeheaver G, et al. Definitions and guidelines for assessment of wounds and evaluation of healing. *Wound Repair Regen* 1994;2(3):165–70.
- [4] Posnett J, Franks PJ. The burden of chronic wounds in the UK. *Nurs Times* 2008;104(3):44–5.
- [5] Kobza L, Scheurich A. The impact of telemedicine on outcomes of chronic wounds in the home care setting. *Ostomy Wound Manage* 2000;46(10):48–53.
- [6] Iyun AO, Ademola SA, Olawoye OA, Michael AI, Oluwatosin OM. Point prevalence of chronic wounds at a tertiary hospital in Nigeria. *Wounds* 2016;28(2):57–62.
- [7] Graves N, Zheng H. The prevalence and incidence of chronic wounds: a literature review. *Wound Pract Res* 2014;22:4–19.
- [8] Sen CK, Gordillo GM, Roy S, Kirsner R, Lambert L, Hunt TK. Human skin wounds: a major and snowballing threat to public health and the economy. *Wound Repair Regen* 2009;17(6):763–71.
- [9] Phillips CJ, Humphreys I, Fletcher J, Harding K, Chamberlain G, Macey S. Estimating the costs associated with the management of patients with chronic wounds using linked routine data. *Int Wound J* 2016;13(6):1193–7.
- [10] Guest JF, Ayoub N, McIlwraith T, Uchegbu I, Gerrish A, Weidlich D, et al. Health economic burden that different wound types impose on the UK's National Health Service. *Int Wound J* 2017;14(2):322–30.
- [11] Fife C, Carter M. Wound care outcomes and associated cost among patients treated in US outpatient wound centers - data from the US wound registry. *Wounds* 2012;24(1):10–7.
- [12] Frykberg RG, Banks J. Challenges in the treatment of chronic wounds. *Adv Wound Care (New Rochelle)* 2015;4(9):560–82.
- [13] Ward MM. Estimating disease prevalence and incidence using administrative data: some assembly required. *J Rheumatol* 2013;40(8):1241–3.
- [14] Wright J, Williams R, Wilkinson JR. Development and importance of health needs assessment. *BMJ* 1998;316(7140):1310–3.
- [15] Divajeva D, Marsh T, Logstrup S, Kestens M, Vemer P, Kriaucioniene V, et al. Economics of chronic diseases protocol: cost-effectiveness modelling and the future burden of non-communicable disease in Europe. *BMC Public Health* 2014;14:456.
- [16] Barendregt JJ, Doi SA, Lee YY, Norman RE, Vos T. Meta-analysis of prevalence. *J Epidemiol Community Health* 2013;67(11):974–8.
- [17] Moher D, Shamseer L, Clarke M, Ghersi D, Liberati A, Petticrew M, et al. Preferred reporting items for systematic review and meta-analysis protocols (PRISMA-P) 2015 statement. *Syst Rev* 2015;4:1.
- [18] Jarbrink K, Ni G, Sönnegren H, Schmidtchen A, Pang C, Bajpai R, et al. Prevalence and incidence of chronic wounds and related complications: a protocol for a systematic review. *Syst Rev* 2016;5(1):152.
- [19] Sheehan P, Jones P, Caselli A, Giurini JM, Veves A. Percent change in wound area of diabetic foot ulcers over a 4-week period is a robust predictor of complete healing in a 12-week prospective trial. *Diabetes Care* 2003;26(6):1879–82.
- [20] van Rijswijk L, Polansky M. Predictors of time to healing deep pressure ulcers. *Ostomy Wound Manage* 1994;40(8):40–2. 46–48 passim.
- [21] Phillips TJ, Machado F, Trout R, Porter J, Olin J, Falanga V. Prognostic indicators in venous ulcers. *J Am Acad Dermatol* 2000;43(4):627–30.
- [22] Munn Z, Moola S, Riitano D, Lisy K. The development of a critical appraisal tool for use in systematic reviews addressing questions of prevalence. *Int J Health Policy Manag* 2014;3:123–8.
- [23] DerSimonian R, Laird N. Meta-analysis in clinical trials. *Control Clin Trials* 1986;7:177–88.
- [24] Clopper CJ, Pearson ES. The use of confidence or fiducial limits illustrated in the case of the binomial. *Biometrika* 1934;26:404–13.
- [25] Higgins JPT, Green S. *Cochrane handbook for systematic reviews of interventions* version 5.1.0 [updated March 2011]. The Cochrane Collaboration; 2011. <http://handbook.cochrane.org>.

- [26] Higgins JP, Thompson SG, Tierney J, Rydzewska L, Burdett S, Stewart L, et al., editors. *Presenting random effects meta-analyses: where are we going wrong*. Lyon, France: International Cochrane Colloquium; 2001.
- [27] Egger M, Smith GD, Schneider M, Minder C. Bias in meta-analysis detected by a simple, graphical test. *BMJ* 1997;315:629–34.
- [28] Baujat B, Mahe C, Pignon JP, Hill C. A graphical method for exploring heterogeneity in meta-analyses: application to a meta-analysis of 65 trials. *Stat Med* 2002;21(18):2641–52.
- [29] Borenstein M, Hedges LV, Higgins JPT, Rothstein HR. *Introduction to meta-analysis*. Chichester, UK: John Wiley & Sons, Ltd; 2009.
- [30] Schwarzer G. meta: An R package for meta-analysis. *R News* 2007;7(3):40–5.
- [31] Heyer K, Herberger K, Protz K, Glaeske G, Augustin M. Epidemiology of chronic wounds in Germany: analysis of statutory health insurance data. *Wound Repair Regen* 2016;24(2):434–42.
- [32] Clarke-Moloney M, Keane N, Kavanagh E. An exploration of current leg ulcer management practices in an Irish community setting. *J Wound Care* 2006;15(9):407–10.
- [33] Moffatt CJ, Franks PJ, Doherty DC, Martin R, Blewett R, Ross F. Prevalence of leg ulceration in a London population. *QJM* 2004;97(7):431–7.
- [34] O'Brien JF, Grace PA, Perry IJ, Burke PE. Prevalence and aetiology of leg ulcers in Ireland. *Ir J Med Sci* 2000;169(2):110–2.
- [35] Oien RF, Hakansson A, Ovred I, Hansen BU. Wound management for 287 patients with chronic leg ulcers demands 12 full-time nurses. Leg ulcer epidemiology and care in a well-defined population in southern Sweden. *Scand J Prim Health Care* 2000;18(4):220–5.
- [36] Oien RF, Ragnarson Tennvall G. Accurate diagnosis and effective treatment of leg ulcers reduce prevalence, care time and costs. *J Wound Care* 2006;15(6):259–62.
- [37] Pálsdóttir G. Chronic leg ulcers among the Icelandic population. *EWMA J* 2010;10(1):19–23. 5.
- [38] Pina E, Furtado K, Franks PJ, Moffatt CJ. Leg ulceration in Portugal: prevalence and clinical history. *Eur J Vasc Endovasc Surg* 2005;29(5):549–53.
- [39] Walker N, Rodgers A, Birchall N, Norton R, MacMahon S. The occurrence of leg ulcers in Auckland: results of a population-based study. *N Z Med J* 2002;115(1151):159–62.
- [40] Hall J, Buckley HL, Lamb KA, Stubbs N, Saramago P, Dumville JC, et al. Point prevalence of complex wounds in a defined United Kingdom population. *Wound Repair Regen* 2014;22(6):694–700.
- [41] McDermott-Scales L, Cowman S, Gethin G. Prevalence of wounds in a community care setting in Ireland. *J Wound Care* 2009;18(10):405–17.
- [42] Jiang Y, Huang S, Fu X, Liu H, Ran X, Lu S, et al. Epidemiology of chronic cutaneous wounds in China. *Wound Repair Regen* 2011;19(2):181–8.
- [43] Rodrigues I, Megie MF. Prevalence of chronic wounds in Quebec home care: an exploratory study. *Ostomy Wound Manage* 2006;52(5):46–8. 52–57.
- [44] Schwien T, Gilbert J, Lang C. Pressure ulcer prevalence and the role of negative pressure wound therapy in home health quality outcomes. *Ostomy Wound Manage* 2005;51(9):47–60.
- [45] Barrois B, Labalette C, Rousseau P, Corbin A, Colin D, Allaert F, et al. A national prevalence study of pressure ulcers in French hospital inpatients. *J Wound Care* 2008;17(9):373–6. 378–379.
- [46] Amir Y, Lohrmann C, Halfens RJ, Schols JM. Pressure ulcers in four Indonesian hospitals: prevalence, patient characteristics, ulcer characteristics, prevention and treatment. *Int Wound J* 2017;14(1):184–93.
- [47] El-Nahas MR, Gawish HMS, Tarshoby MM, State OI, Boulton AJM. The prevalence of risk factors for foot ulceration in Egyptian diabetic patients. *Pract Diabetes Int* 2008;25(9):362–6. 5.
- [48] Nelzen O. Prevalence of venous leg ulcer: the importance of the data collection method. *Phlebology* 2008;15(4):143–50.
- [49] Fonder MA, Lazarus GS, Cowan DA, Aronson-Cook B, Kohli AR, Mamelak AJ. Treating the chronic wound: a practical approach to the care of nonhealing wounds and wound care dressings. *J Am Acad Dermatol* 2008;58(2):185–206.
- [50] Baker SR, Stacey MC. Epidemiology of chronic leg ulcers in Australia. *Aust N Z J Surg* 1994;64(4):258–61.
- [51] Gould L, Abadir P, Brem H, Carter M, Conner-Kerr T, Davidson J, et al. Chronic wound repair and healing in older adults: current status and future research. *Wound Repair Regen* 2015;23(1):1–13.
- [52] Gupta N, Gupta SK, Shukla VK, Singh SP. An Indian community-based epidemiological study of wounds. *J Wound Care* 2004;13(8):323–5.
- [53] Cho NH, Shaw JE, Karuranga S, Huang Y, da Rocha Fernandes JD, Ohlrogge AW, et al. IDF Diabetes Atlas: global estimates of diabetes prevalence for 2017 and projections for 2045. *Diabetes Res Clin Pract* 2018;138:271–81.
- [54] Pearce N. Global epidemiology: the importance of international comparisons and collaborations. *Open Access Epidemiol* 2013;1(2):15.
- [55] Ledikwe JH, Grignon J, Lebelonyane R, Ludick S, Matshediso E, Sento BW, et al. Improving the quality of health information: a qualitative assessment of data management and reporting systems in Botswana. *Health Res Pol Syst* 2014;12:7.
- [56] Frerichs RR. Epidemiologic surveillance in developing countries. *Annu Rev Public Health* 1991;12:257–80.
- [57] Gupta SK, Shukla VK. Leg ulcers in the tropics. *Int J Low Extrem Wounds* 2002;1(1):58–61.
- [58] Karimkhani C, Dellavalle RP, Coffeng LE, Flohr C, Hay RJ, Langan SM, et al. Global skin disease morbidity and mortality: an update from the global burden of disease study 2013. *JAMA Dermatol* 2017;153(5):406–12.
- [59] Hay RJ, Johns NE, Williams HC, Bolliger IW, Dellavalle RP, Margolis DJ, et al. The global burden of skin disease in 2010: an analysis of the prevalence and impact of skin conditions. *J Invest Dermatol* 2014;134(6):1527–34.
- [60] Mostow EN. Diagnosis and classification of chronic wounds. *Clin Dermatol* 1994;12(1):3–9.